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<110> Intellectual Property Consulting, Inc. TOHYAMA, Masaya YAMASHITA, Toshihide TANAKA, Hiroyuki HIGUCHI, Haruhisa

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Asp Leu Glu Glu Leu Glu Val Leu Glu Arg Lys Pro Ala Ala Gly Leu 50 55 60

Ser Ala Ala Pro Val Pro Pro Ala Ala Ala Pro Leu Leu Asp Phe Ser 65 70 75 80

Ser Asp Ser Val Pro Pro Ala Pro Arg Gly Pro Leu Pro Ala Ala Pro 85 90 95

Pro Thr Ala Pro Glu Arg Gln Pro Ser Trp Glu Arg Ser Pro Ala Ala 100 105 110 Ser Ala Pro Ser Leu Pro Pro Ala Ala Ala Val Leu Pro Ser Lys Leu 115 120 125

Pro Glu Asp Asp Glu Pro Pro Ala Arg Pro Pro Ala Pro Ala Gly Ala

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Ser Pro Leu Ala Glu Pro Ala Ala Pro Pro Ser Thr Pro Ala Ala Pro 145 150 155 160

Lys Arg Arg Gly Ser Gly Ser Val Asp Glu Thr Leu Phe Ala Leu Pro 165 170 175

Ala Ala Ser Glu Pro Val IIe Pro Ser Ser Ala Glu Lys IIe Met Asp 180 185 190

Leu Lys Glu Gln Pro Gly Asn Thr Val Ser Ser Gly Gln Glu Asp Phe 195 200 205

Pro Ser Val Leu Phe Glu Thr Ala Ala Ser Leu Pro Ser Leu Ser Pro 210 215 220

Leu Ser Thr Val Ser Phe Lys Glu His Gly Tyr Leu Gly Asn Leu Ser 225 230 235 240

Ala Val Ala Ser Thr Glu Gly Thr lie Glu Glu Thr Leu Asn Glu Ala 245 250 255 Ser Arg Glu Leu Pro Glu Arg Ala Thr Asn Pro Phe Val Asn Arg Glu 260 265 270

Ser Ala Glu Phe Ser Val Leu Glu Tyr Ser Glu Met Gly Ser Ser Phe 275 280 285

Asn Gly Ser Pro Lys Gly Glu Ser Ala Met Leu Val Glu Asn Thr Lys 290 295 300

Glu Glu Val IIe Val Arg Ser Lys Asp Lys Glu Asp Leu Val Cys Ser 305 310 315 320

Ala Ala Leu His Asn Pro Gln Glu Ser Pro Ala Thr Leu Thr Lys Val 325 330 335

Val Lys Glu Asp Gly Val Met Ser Pro Glu Lys Thr Met Asp 11e Phe 340 345 350

Asn Giu Met Lys Met Ser Val Val Ala Pro Val Arg Giu Giu Tyr Ala 355 360 365

Asp Phe Lys Pro Phe Glu Gln Ala Trp Glu Val Lys Asp Thr Tyr Glu 370 380 .

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Leu Val Lys Asp Gly Ser Arg Ala Tyr IIe Thr Cys Asp Ser Phe Ser 435 440 445

Ser Ala Thr Glu Ser Thr Ala Ala Asn ile Phe Pro Val Leu Glu Asp 450 455 460

His Thr Ser Glu Asn Lys Thr Asp Glu Lys Lys IIe Glu Glu Arg Lys
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Ala Gin ile ile Thr Giu Lys Thr Ser Pro Lys Thr Ser Asn Pro Phe
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Leu Val Ala IIe His Asp Ser Glu Ala Asp Tyr Val Thr Thr Asp Asn 500 505 510

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Ser Glu Ala IIe Gln Glu Ser IIe Tyr Pro Thr Ala Gln Leu Cys Pro 565 570 575

Ser Phe Glu Glu Ala Glu Ala Thr Pro Ser Pro Val Leu Pro Asp lle 580 585 590

Val Met Glu Ala Pro Leu Asn Ser Leu Leu Pro Ser Thr Gly Ala Ser 595 600 605

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Tyr Asp Gly IIe Lys Leu Glu Pro Glu Asn Pro Pro Pro Tyr Glu Glu 625 630 635 640

Ala Met Ser Val Ala Leu Lys Thr Ser Asp Ser Lys Glu Glu lle Lys . 645 650 655

Glu Pro Glu Ser Phe Asn Ala Ala Ala Gln Glu Ala Glu Ala Pro Tyr 660 665 670

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Glu Pro Val Asp Leu Phe Ser Asp Asp Ser IIe Pro Glu Val Pro Gln
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Ser Glu Thr Val Thr Gln His Lys His Lys Glu Arg Leu Ser Ala Ser 755 760 765

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Glu Val Ser Asn Lys Ser Glu ile Ala Asn Val Gin Ser Gly Ala Asn 865 870 875 880

Ser Leu Pro Cys Ser Glu Leu Pro Cys Asp Leu Ser Phe Lys Asn Thr 885 890 895

Tyr Pro Lys Asp Glu Ala His Val Ser Asp Glu Phe Ser Lys Ser Arg 900 905 910

Ser Ser Val Ser Lys Val Pro Leu Leu Leu Pro Asn Val Ser Ala Leu 915 920 925

Glu Ser Gin He Glu Met Gly Asn He Val Lys Pro Lys Val Leu Thr 930 935 940

Lys Glu Ala Glu Glu Lys Leu Pro Ser Asp Thr Glu Lys Glu Asp Arg 945 950 955 960

Ser Leu Thr Ala Val Leu Ser Ala Glu Leu Asn Lys Thr Ser Val Val 965 970 975 Asp Leu Leu Tyr Trp Arg Asp ile Lys Lys Thr Gly Val Val Phe Gly 980 985 990

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Val Thr Ala Tyr lle Ala Leu Ala Leu Leu Ser Val Thr lle Ser 1010 1015 1020

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Glu Gly His Pro Phe Arg Ala Tyr Leu Glu Ser Glu Val Ala lle 1040 1045 1050

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Gln Ala Gln IIe Asp His Tyr Leu Gly Leu Ala Asn Lys Ser Val 1130 1135 1140

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20 25 30

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Asp Tyr Asp Arg Leu Arg Pro Leu Ser Tyr Pro Asp Thr Asp Val IIe 65 70 75 80 Leu Met Cys Phe Ser IIe Asp Ser Pro Asp Ser Leu Glu Asn IIe Pro 85 90 95

Glu Lys Trp Thr Pro Glu Val Lys His Phe Cys Pro Asn Val Pro IIe 100 105 110

Ile Leu Val Gly Asn Lys Lys Asp Leu Arg Asn Asp Glu His Thr Arg 115 120 125

Arg Glu Leu Ala Lys Met Lys Gin Glu Pro Vai Lys Pro Glu Glu Gly
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Arg Asp Met Ala Asn Arg IIe Gly Ala Phe Gly Tyr Met Glu Cys Ser 145 150 155 160

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Asn Phe Asp Phe Val Thr Glu Thr Pro Leu Glu Gly Asn Phe Val Trp 50 55 60

Glu Arg Val Arg Ser Leu Gly Leu Pro Lys Val Tyr Leu Ser Pro Gly
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Ser Arg Ser Arg Asp Leu Gly Gly Asp Lys Arg Pro Ser Thr Ser 85 90 95 Ser Ala Leu Gin Gly Pro Ala Pro Glu Asp His Val Ala Leu Ser 100 105 110

Leu Ser Cys Thr Leu Val Ser Glu Arg Pro Glu Asp Ser Pro Gly Gly
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Pro Gly Thr Ser Gln Gly Arg Lys Arg Arg Gln Thr Ser Leu Thr Asp 130 135 140

Phe Tyr His Ser Lys Arg Arg Leu Val Phe Cys Lys Arg Lys Pro 145 150 155

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<212> PRT

<213> Rattus norvegicus

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Cys Ser Thr Gly Leu Tyr Thr His Ser Gly Glu Cys Cys Lys Ala Cys 35 40 45

Asn Leu Gly Glu Gly Val Ala Gin Pro Cys Gly Ala Asn Gin Thr Val
50 55 60

Cys Glu Pro Cys Leu Asp Asn Val Thr Phe Ser Asp Val Val Ser Ala 65 70 75 80

Thr Glu Pro Cys Lys Pro Cys Thr Glu Cys Leu Gly Leu Gln Ser Met
85 90 95

Ser Ala Pro Cys Val Glu Ala Asp Asp Ala Val Cys Arg Cys Ala Tyr 100 105 110

Gly Tyr Tyr Gln Asp Glu Glu Thr Gly His Cys Glu Ala Cys Ser Val

Cys Glu Val Gly Ser Gly Leu Val Phe Ser Cys Gin Asp Lys Gin Asn 130 135 140

Thr Val Cys Glu Glu Cys Pro Glu Gly Thr Tyr Ser Asp Glu Ala Asn 145 150 155 160

His Val Asp Pro Cys Leu Pro Cys Thr Val Cys Glu Asp Thr Glu Arg 165 170 175

Gin Leu Arg Giu Cys Thr Pro Trp Ala Asp Ala Giu Cys Giu Giu Ile 180 185 190

Pro Gly Arg Trp lie Pro Arg Ser Thr Pro Pro Glu Gly Ser Asp Ser 195 200 205

Thr Ala Pro Ser Thr Gln Glu Pro Glu Val Pro Pro Glu Gln Asp Leu 210 215 220

Val Pro Ser Thr Val Ala Asp Met Val Thr Thr Val Met Gly Ser Ser 225 230 235 240

Gin Pro Val Val Thr Arg Gly Thr Thr Asp Asn Leu IIe Pro Val Tyr
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Phe Lys Arg Trp Asn Ser Cys Lys Gln Asn Lys Gln Gly Ala Asn Ser 275 280 285

Arg Pro Val Asn Gln Thr Pro Pro Pro Glu Gly Glu Lys Leu His Ser 290 295 300

Asp Ser Gly IIe Ser Val Asp Ser Gln Ser Leu His Asp Gln Gln Thr 305 310 315 320

His Thr Gln Thr Ala Ser Gly Gln Ala Leu Lys Gly Asp Gly Asn Leu 325 330 335

Tyr Ser Ser Leu Pro Leu Thr Lys Arg Glu Glu Val Glu Lys Leu Leu 340 345 350

Asn Gly Asp Thr Trp Arg His Leu Ala Gly Glu Leu Gly Tyr Gln Pro 355 360 365

Glu His Ile Asp Ser Phe Thr His Glu Ala Cys Pro Val Arg Ala Leu 370 375 380 Leu Ala Ser Trp Gly Ala Gln Asp Ser Ala Thr Leu Asp Ala Leu Leu 385 390 395 400

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<211> 4167

<212> DNA

<213> Homo sapiens

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<213> Homo sapiens

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Gly Leu Asn Ser Leu Val Leu Asp Leu Asp Phe Pro Ala Leu Arg Lys
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Asn Lys Asn IIe Asp Asn Phe Leu Asn Arg Tyr Glu Lys IIe Val Lys 65 70 75 80

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Val 11e Gly Arg Gly Ala Phe Gly Glu Val Gln Leu Val Arg His Lys 100 105 110

Ala Ser Gin Lys Val Tyr Ala Met Lys Leu Leu Ser Lys Phe Glu Met 115 120 125

Ala Phe Ala Asn Ser Pro Trp Val Val Gin Leu Phe Tyr Ala Phe Gin 145 150 155 160

Asp Asp Arg Tyr Leu Tyr Met Val Met Glu Tyr Met Pro Gly Gly Asp 165 170 175

Leu Val Asn Leu Met Ser Asn Tyr Asp Val Pro Glu Lys Trp Ala Lys 180 185 190

Phe Tyr Thr Ala Glu Val Val Leu Ala Leu Asp Ala IIe His Ser Met
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Gly Leu lle His Arg Asp Val Lys Pro Asp Asn Met Leu Leu Asp Lys 210 215 220

His Gly His Leu Lys Leu Ala Asp Phe Gly Thr Cys Met Lys Met Asp 225 230 235 240

Glu Thr Gly Met Val His Cys Asp Thr Ala Val Gly Thr Pro Asp Tyr
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lle Ser Pro Glu Val Leu Lys Ser Gln Gly Gly Asp Gly Phe Tyr Gly 260 265 270

Arg Glu Cys Asp Trp Trp Ser Val Gly Val Phe Leu Tyr Glu Met Leu 275 280 285

Val Gly Asp Thr Pro Phe Tyr Ala Asp Ser Leu Val Gly Thr Tyr Ser 290 295 300

Lys IIe Met Asp His Lys Asn Ser Leu Cys Phe Pro Glu Asp Ala Glu 305 310 315 320

lle Ser Lys His Ala Lys Asn Leu lle Cys Ala Phe Leu Thr Asp Arg 325 330 335

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Asp Asp IIe Glu Asp Asp Lys Gly Asp Val Glu Thr Phe Pro IIe Pro 385 390 395 400

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Glu Leu Glu Gin Lys Cys Lys Ser Val Asn Thr Arg Leu Glu Lys Thr 465 470 475 480

Ala Lys Glu Leu Glu Glu Glu IIe Thr Leu Arg Lys Ser Val Glu Ser 485 490 495 Ala Leu Arg Gln Leu Glu Arg Glu Lys Ala Leu Leu Gln His Lys Asn 500 505 510

Ala Glu Tyr Gln Arg Lys Ala Asp His Glu Ala Asp Lys Lys Arg Asn 515 520 525

Leu Glu Asn Asp Val Asn Ser Leu Lys Asp Gln Leu Glu Asp Leu Lys 530 535 540

Lys Arg Asn Gln Asn Ser Gln IIe Ser Thr Glu Lys Val Asn Gln Leu 545 550 555 560

Gin Arg Gin Leu Asp Giu Thr Asn Ala Leu Leu Arg Thr Giu Ser Asp 565 570 575

Thr Ala Ala Arg Leu Arg Lys Thr Gln Ala Glu Ser Ser Lys Gln IIe 580 585 590

Gin Gin Leu Giu Ser Asn Asn Arg Asp Leu Gin Asp Lys Asn Cys Leu 595 600 605

Leu Glu Thr Ala Lys Leu Lys Leu Glu Lys Glu Phe lle Asn Leu Gln 610 615 620

Ser Ala Leu Glu Ser Glu Arg Arg Asp Arg Thr His Gly Ser Glu lle 625 630 635 640 lle Asn Asp Leu Gin Gly Arg IIe Cys Gly Leu Glu Glu Asp Leu Lys
645 650 655

Asn Gly Lys lle Leu Leu Ala Lys Val Glu Leu Glu Lys Arg Gln Leu 660 665 670

Gin Glu Arg Phe Thr Asp Leu Glu Lys Glu Lys Ser Asn Met Glu Ile[.]
675 680 685

Asp Met Thr Tyr Gin Leu Lys Val lie Gin Gin Ser Leu Giu Gin Giu 690 695 700

Glu Ala Glu His Lys Ala Thr Lys Ala Arg Leu Ala Asp Lys Asn Lys 705 710 715 720

lle Tyr Glu Ser ile Glu Glu Ala Lys Ser Glu Ala Met Lys Glu Met 725 730 735

Glu Lys Lys Leu Leu Glu Glu Arg Thr Leu Lys Gln Lys Val Glu Asn 740 745 750

Leu Leu Glu Ala Glu Lys Arg Cys Ser Leu Leu Asp Cys Asp Leu 755 760 765

Lys Gin Ser Gin Gin Lys IIe Asn Giu Leu Leu Lys Gin Lys Asp Val 770 775 780 Leu Asn Glu Asp Val Arg Asn Leu Thr Leu Lys lle Glu Gln Glu Thr 785 790 795 800

Gin Lys Arg Cys Leu Thr Gin Asn Asp Leu Lys Met Gin Thr Gin Gin 805 810 815

Val Asn Thr Leu Lys Met Ser Glu Lys Gln Leu Lys Gln Glu Asn Asn 820 825 830

His Leu Met Glu Met Lys Met Asn Leu Glu Lys Gin Asn Ala Glu Leu 835 840 845

Arg Lys Glu Arg Gln Asp Ala Asp Gly Gln Met Lys Glu Leu Gln Asp 850 855 860

Gin Leu Giu Ala Giu Gin Tyr Phe Ser Thr Leu Tyr Lys Thr Gin Vai 865 870 875 880

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Leu Gin Gin Lys Lys Gin Glu Leu Gin Asp Glu Arg Asp Ser Leu Ala 900 905 910

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Met Lys Glu Leu Glu lle Lys Glu Met Met Ala Arg His Lys Gln Glu 945 955 960

Leu Thr Glu Lys Asp Ala Thr IIe Ala Ser Leu Glu Glu Thr Asn Arg 965 970 975

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Val Gly Glu Lys Ser Asn Tyr I le Cys His Lys Gly His Glu Phe 1250 1255 1260

Ile Pro Thr Leu Tyr His Phe Pro Thr Asn Cys Glu Ala Cys Met 1265 1270 1275

Lys Pro Leu Trp His Met Phe Lys Pro Pro Pro Ala Leu Glu Cys 1280 1285 1290

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Asp Cys Asp Ala Leu Met Ala Gly Cys lle Gln Glu Ala Arg Glu Arg 35 40 45

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Trp Glu Arg Val Arg Gly Leu Gly Leu Pro Lys Leu Tyr Leu Pro Thr 65 70 75 80

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Thr Ser Pro Ala Leu Leu Gin Gly Thr Ala Glu Glu Asp His Val Asp 100 105 110

Leu Ser Leu Ser Cys Thr Leu Val Pro Arg Ser Gly Glu Gln Ala Glu 115 120 125

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Phe Cys Ser His Cys Thr Asp Phe IIe Trp Gly Phe Gly Lys Gln Gly 50 55 60

Phe Gln Cys Gln Val Cys Cys Phe Val Val His Lys Arg Cys His Glu 65 70 75 80

Phe Val Thr Phe Ser Cys Pro Gly Ala Asp Lys Gly Pro Asp Thr Asp 85 90 95

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Arg lie Tyr Leu Lys Ala Glu Val Thr Asp Glu Lys Leu His Val Thr 165 170 175

Val Arg Asp Ala Lys Asn Leu lie Pro Met Asp Pro Asn Gly Leu Ser

180

185

190

Asp Pro Tyr Val Lys Leu Lys Leu IIe Pro Asp Pro Lys Asn Glu Ser 195 200 205

Lys Gin Lys Thr Lys Thr lie Arg Ser Thr Leu Asn Pro Gin Trp Asn 210 215 220

Glu Ser Phe Thr Phe Lys Leu Lys Pro Ser Asp Lys Asp Arg Arg Leu 225 230 235 240

Ser Val Glu lle Trp Asp Trp Asp Arg Thr Thr Arg Asn Asp Phe Met 245 250 255

Gly Ser Leu Ser Phe Gly Val Ser Glu Leu Met Lys Met Pro Ala Ser 260 265 270

Gly Trp Tyr Lys Leu Leu Asn Gin Glu Glu Glu Glu Tyr Tyr Asn Val 275 280 285

Pro Ile Pro Glu Gly Asp Glu Glu Gly Asn Val Glu Leu Arg Gln Lys 290 295 300

Phe Glu Lys Ala Lys Leu Gly Pro Ala Gly Asn Lys Val IIe Ser Pro 305 310 315 320

Ser Glu Asp Arg Lys Gln Pro Ser Asn Asn Leu Asp Arg Val Lys Leu

325 .

330

335

Thr Asp Phe Asn Phe Leu Met Val Leu Gly Lys Gly Ser Phe Gly Lys 340 345 350

Val Met Leu Ala Asp Arg Lys Gly Thr Glu Glu Leu Tyr Ala lle Lys 355 360 365

ile Leu Lys Lys Asp Val Val lie Gin Asp Asp Val Giu Cys Thr 370 375 380

Met Val Glu Lys Arg Val Leu Ala Leu Leu Asp Lys Pro Pro Phe Leu 385 390 395 400

Thr Gln Leu His Ser Cys Phe Gln Thr Val Asp Arg Leu Tyr Phe Val
405 410 415

Met Glu Tyr Val Asn Gly Gly Asp Leu Met Tyr His ile Gln Gln Val 420 425 430

Gly Lys Phe Lys Glu Pro Gln Ala Val Phe Tyr Ala Ala Glu Ile Ser 435 440 445

lle Gly Leu Phe Phe Leu His Lys Arg Gly lle lle Tyr Arg Asp Leu 450 455 460

Lys Leu Asp Asn Val Met Leu Asp Ser Glu Gly His lle Lys lle Ala

480

465 470 475

Asp Phe Gly Met Cys Lys Glu His Met Met Asp Gly Val Thr Thr Arg
485 490 495

Thr Phe Cys Gly Thr Pro Asp Tyr lie Ala Pro Glu lie lie Ala Tyr
500 505 510

Gin Pro Tyr Gly Lys Ser Val Asp Trp Trp Ala Tyr Gly Val Leu Leu 515 520 525

Tyr Glu Met Leu Ala Gly Gln Pro Pro Phe Asp Gly Glu Asp Glu Asp 530 535 540

Glu Leu Phe Gln Ser Ile Met Glu His Asn Val Ser Tyr Pro Lys Ser 545 550 555 560

Leu Ser Lys Glu Ala Val Ser lle Cys Lys Gly Leu Met Thr Lys His
565 570 575.

Pro Ala Lys Arg Leu Gly Cys Gly Pro Glu Gly Glu Arg Asp Val Arg 580 585 590

Glu His Ala Phe Phe Arg Arg Ile Asp Trp Glu Lys Leu Glu Asn Arg 595 600 605

Glu lle Gln Pro Pro Phe Lys Pro Lys Val Cys Gly Lys Gly Ala Glu

610 615 620

Asn Phe Asp Lys Phe Phe Thr Arg Gly Gln Pro Val Leu Thr Pro Pro 625 630 635 640

Asp Gln Leu Val lie Ala Asn lie Asp Gln Ser Asp Phe Glu Gly Phe 645 650 655

Ser Tyr Val Asn Pro Gln Phe Val His Pro 11e Leu Gln Ser Ala Val 660 665 670